A CONFIRMATION OF THE OLIGODENDROGLIAL ORIGIN OF MYELIN IN THE ADULT RAT

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It is widely held that myelin sheaths in the central nervous system are formed by an oligodendroglial process which wraps itself around an axon (1-5, 9-21). We have, in our own work, assumed this to be the case and, indeed, have presented some indirect evidence that the myelin sheath was, at least at times, connected to a glial perikaryon. This evidence consisted of the presence of formed organelles such as mitochondria and dense bodies within the outer, inner, and lateral loops of the myelin sheath (6-8). In addition, we interpreted the presence of unusually long outer loops as indicating a probable connection to a glial perikaryon

(8). Furthermore, it is well known that the cytoplasmic portions of the myelin sheath greatly resemble the cytoplasm of the oligodendroglial cell.

A distinct connection has been convincingly demonstrated in the young, developing animal by Bunge et al. (2) and by Peters (17). In the adult animal, however, a connection between a mature myelin sheath and a glial cell has not been clearly demonstrated. Only a single micrograph occurs in the literature which purports to demonstrate this connection (Fig. 6 in Kruger and Maxwell, reference 9). If this is the only direct demonstra-

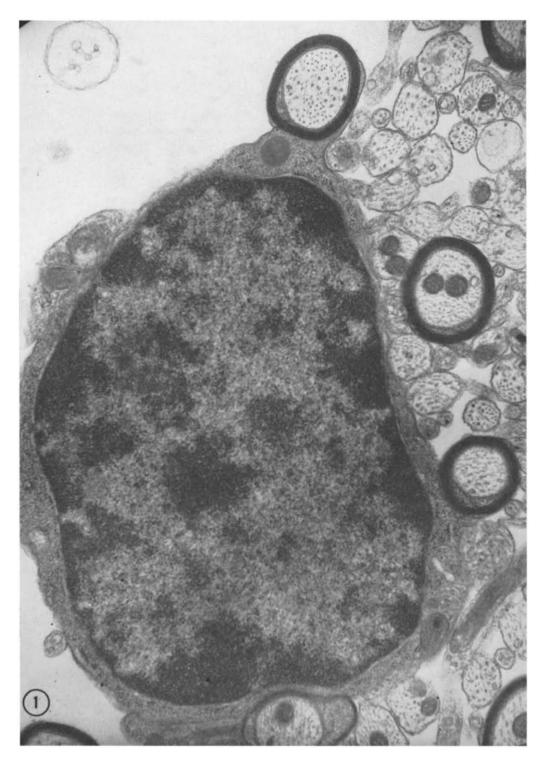


FIGURE 1 An oligodendroglial cell at the border of a cystic space in the white matter of an adult rat cerebrum. An apparently mature myelin sheath whose outermost lamella appears continuous with the perikaryon of the oligodendroglial cell is visible in the upper portion of the micrograph. \times 25,000.

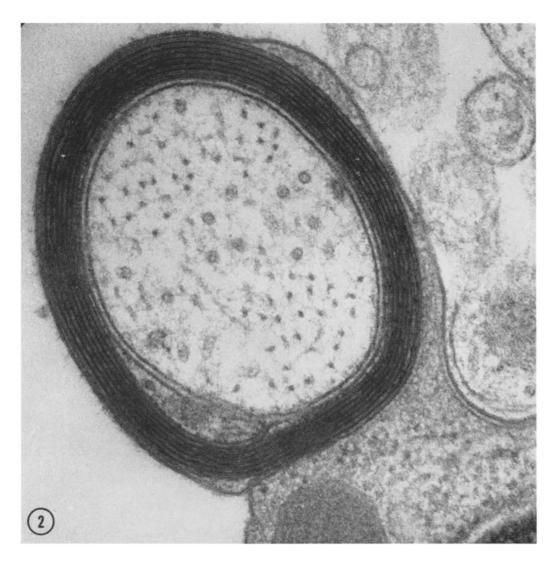


FIGURE 2 A higher magnification of a portion of Fig. 1 showing the continuity between the outermost lamella and the oligodendroglial plasma membrane. \times 110,000.

tion of a glial connection to the myelin sheath in the adult animal, as recently stated in an extensive review by Bunge (1), then the evidence for this phenomenon is indeed shaky.

During an experimental study of central white matter in the adult rat, we happened to come across a section which, to our mind, convincingly demonstrates such a direct connection between an apparently mature myelin sheath and an oligodendrocyte (Figs. 1, 2). This material, which had been prepared for electron microscopy by techniques previously described (6–8) was obtained from the edge of a cyst which was formed in the brain of an adult rat 3 wk after the intracerebral implantation of a pellet of triethyltin acetate.

The myelin sheath is apparently mature, but one must bear in mind that it is in an area in which tissue damage had previously occurred. Therefore, one may not exclude the possibility that this configuration is, in fact, a late stage of remyelination. If so, the micrograph offers excellent confirmation of remyelination within the central nervous system. If not, then we have finally, due to a purely fortuitous section, found a direct connection between a mature myelin sheath and an oligodendrocyte in the adult central nervous system.

This investigation was supported by United States Public Health Service research grant No. B-3533

REFERENCES

- 1. BUNGE, R. P. 1968. Glial cells and the central myelin sheath. *Physiol. Rev.* 48:197.
- BUNGE, M. B., R. P. BUNGE, and G. D. PAPPAS. 1962. Electron microscopic demonstration of connections between glia and myelin sheaths in the developing mammalian central nervous system. J. Cell Biol. 12:448.
- BUNGE, M. B., R. P. BUNGE, and H. RIS. 1961. Ultrastructural study of remyelination in an experimental lesion in adult cat spinal cord. J. Biophys. Biochem. Cytol. 10:67.
- BUNGE, R. P., and P. M. GLASS. 1965. Some observations on myelin-glial relationships and on the etiology of the cerebrospinal fluid exchange lesion. Ann. N.Y. Acad. Sci. 122:15.
- HERNDON, R. M. 1964. The fine structure of the rat cerebellum. II. The stellate neurons, granule cells, and glia. J. Cell Biol. 23:277.
- HIRANO, A., and H. M. DEMBITZER. 1967. A structural analysis of the myelin sheath in the central nervous system. J. Cell Biol. 34:555.
- HIRANO, A., S. LEVINE, and H. M. ZIMMERMAN. 1968. Remyelination in the central nervous system after cyanide intoxication. J. Neuropathol. Exptl. Neurol. 27:234.
- HIRANO, A., H. M. ZIMMERMAN, and S. LEVINE. 1966. Myelin in the central nervous system as observed in experimentally induced edema in the rat. J. Cell Biol. 31:397.
- KRUGER, L., and D. S. MAXWELL 1966. Electron microscopy of oligodendrocytes in normal rat cerebrum. Am. J. Anat. 118:411.
- LAMPERT, P. W. 1965. Demyelination and remyelination in experimental allergic encephalomyelitis. Further electron microscopic observations. J. Neuropathol. Exptl. Neurol. 24:371.

from the National Institute of Neurological Diseases and Blindness, National Institutes of Health.

Received for publication 13 March 1968.

- MATURANA, H. R. 1960. The fine anatomy of the optic nerve of Anurans-an electron microscope study. J. Biophys. Biochem. Cytol, 7:107.
- 12. METUZALS, J. 1963. Ultrastructure of myelinated nerve fibers in the central nervous system of the frog. J. Ultrastruct. Res. 8:30.
- METUZALS, J. 1965. The relationship between the glia-satellite cell and the axon in the central myclinated fibers. Proc. European Reg. Conf. Electron Microscopy, 3rd Prague 1964. B:301.
- PETERS, A. 1960. The structure of myelin sheaths in the central nervous system of *Xenopus lavis* (Daudin). J. Biophys. Biochem. Cytol. 7:121.
- PETERS, A. 1960. The formation and structure of myelin sheaths in the central nervous system. J. Biophys. Biochem. Cytol. 8:431.
- PETERS, A. 1964. Further observations on the structure of myelin sheaths in the central nervous system. J. Cell Biol. 20:281.
- PETERS, A. 1964. Observations on the connexions between myelin sheaths and glial cells in the optic nerves of young rats. J. Anat. 98:125.
- PETERS, A. 1966. The node of Ranvier in the central nervous system. *Quart. J. Exptl. Physiol.* 51:229.
- Ross, L. L., M. B. BORNSTEIN, and G. M. LEHRER. 1962. Electron microscopic observations of rat and mouse cerebellum in tissue culture. J. Cell Biol. 14:19.
- UZMAN, B. G. 1964. The spiral configuration of myelin lamellae. J. Ultrastruct. Res. 11:208.
- UZMAN, B. G., and G. M. VILLEGAS. 1960. A comparison of nodes of Ranvier in sciatic nerves with node-like structures in optic nerves of the mouse. J. Biophys. Biochem. Cytol. 7:761.